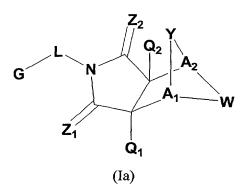
#### Claims

We claim:

#### 1. A compound of the following formula:

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wherein the symbols have the following meanings and are, for each occurrence,

10 independently selected:

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O, S, NH, or NR<sup>6</sup>;

 $Z_2$  is O, S, NH, or NR<sup>6</sup>;

15  $A_1$  is  $CR^7$  or N;

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 $A_2$  is  $CR^7$  or N;

Y' is J-J'-J" where J is  $(CR^7R^{7'})n$  and n=0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, CR<sup>7</sup>R<sup>7'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, or heterocyclo or substituted heterocyclo, and J" is  $(CR^7R^{7'})n$  and n=0-3, where Y is not a bond; and

W' is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein, when W' is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, or heterocyclo or

substituted heterocyclo, then J' must be O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N; or alternatively,

Y' is NR<sup>7</sup>-CR<sup>7</sup>R<sup>7</sup> and W' is CR<sup>8</sup>=CR<sup>8</sup>; or, alternatively,

Y' is  $CR^7R^7$ -C=O and W' is  $NR^9$ - $CR^7R^7$ ;

5 Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R6NC=O,

HOCR<sup>7</sup>R<sup>7</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup> or NR<sup>4</sup>R<sup>5</sup>; 10

O<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R6NC=O, HOCR<sup>7</sup>R<sup>7'</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup> or NR<sup>4</sup>R<sup>5</sup>;

L is a bond,  $(CR^7R^{7'})n$ , NH, NR<sup>5</sup> or NR<sup>5</sup> $(CR^7R^{7'})n$ , where n = 0-3;

- R<sup>1</sup> and R<sup>1</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo 20 or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or 25 substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R<sup>3</sup> and R<sup>3'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or 30 substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl,

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cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, hydroxylamine, hydroxamide, alkoxy or substituted alkoxy, amino, NR<sup>1</sup>R<sup>2</sup>, thiol, alkylthio or substituted alkylthio;

- R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O,
   SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>';
  - R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>:
- R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
- R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkylalkyl or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>3</sub>NR<sup>1</sup>R<sup>1'</sup>, or, wherein

 $A_1$  or  $A_2$  contains a group  $R^7$  and W contains a group  $R^7$ , said  $R^7$  groups of  $A_1$  or  $A_2$  and W together form a heterocyclic ring;

R<sup>8</sup> and R<sup>8'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR<sup>1</sup>, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, alkylthio or substituted alkylthio, C=OSR<sup>1</sup>, R<sup>1</sup>OC=O, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, R<sup>1</sup>R<sup>1'</sup>NC=O, SO<sub>2</sub>OR<sup>1</sup>, S=OR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>; and

R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

with the provisos that:

(1) when Y' is -O-, Q<sub>1</sub> and Q<sub>2</sub> are hydrogen, Z<sub>1</sub> and Z<sub>2</sub> are O, W' is -CH<sub>2</sub>-CH<sub>2</sub>-, and A<sub>1</sub> and A<sub>2</sub> are CH, then G-L is not phenyl, monosubstituted phenyl or phenyl which is substituted with two or more of the following groups: methoxy, halo, NO<sub>2</sub>, methyl, CH<sub>3</sub>-S-, OH, CO<sub>2</sub>H, trifluoromethyl, -C(O)-C<sub>6</sub>H<sub>5</sub>, NH<sub>2</sub>, 4-7-epoxy, hexahydro-1H-isoindole-1,3(2H)dione, or -C(O)-CH<sub>3</sub>;
(2) when Y' is -O-, Q<sub>1</sub> and Q<sub>2</sub> are hydrogen, Z<sub>1</sub> and Z<sub>2</sub> are O, W' is CH<sub>2</sub>-CH<sub>2</sub>, and one of A<sub>1</sub> and A<sub>2</sub> is CH and the other is CR<sup>7</sup>, then G-L is not unsubstituted phenyl;

(3) when Y' is -O-,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is  $CH_2$ - $CH_2$ , and one of  $A_1$  and  $A_2$  is CH and the other is C-CH<sub>3</sub>, then G-L is not phenyl substituted with chloro and/or methyl;

(4) when Y' is -O- or -S-,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is  $CH_2$ -CH<sub>2</sub>, and one of A<sub>1</sub> and A<sub>2</sub> is CH and the other is CH or C-alkyl, then G-L is not N-substituted piperazine-alkyl- or N-substituted imidazolidine-alkyl-; (5) when Y' is -O-;  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is  $CH_2$ - $CH_2$ , and A<sub>1</sub> and A<sub>2</sub> are CH, then G-L is not oxazole or triazole; 5 (6) when Y' is -O-;  $Q_1$  and  $Q_2$  are hydrogen or methyl,  $Z_1$  and  $Z_2$  are O, W' is CH<sub>2</sub>-CH<sub>2</sub>, and A<sub>1</sub> and A<sub>2</sub> are CH or C-CH<sub>3</sub>, then G-L is not thiazole or substituted thiazole; (7) when Y' contains a group J' selected from S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHR<sup>6</sup>, 10 NR<sup>6</sup>NH or N=N, W' is  $CR^7R^7 - CR^7R^7$ , and  $Z_1$  and  $Z_2$  are O, then G-L is not unsubstituted phenyl; (8) when Y is  $NR^7$ , W' is unsubstituted or substituted phenyl, and  $Q_1$  and  $Q_2$ are hydrogen, then  $Z_1$  and  $Z_2$  are not O; (9) when Y' is -O,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is 15 dihydroisoxazole bearing an optionally substituted phenyl group, and A1 and A2 are CH, then G-L is not unsubstituted phenyl or dichlorophenyl; (10) when Y' is O,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is ethylene oxide, and A1 and A2 are CH, then G-L is not methylphenyl or chlorophenyl; (11) when Y' is  $NR^7$ — $CR^7R^7$ , W' is  $CR^8$ = $CR^8$ ,  $Q_1$  and  $Q_2$  are hydrogen,  $A_1$ 20 and  $A_2$  are CH, C-CH<sub>3</sub>, C-CH<sub>2</sub>-C<sub>6</sub>H<sub>5</sub> or C-CH<sub>2</sub>-CH<sub>3</sub>, and  $Z_1$  and  $Z_2$  are O, then G-L is not unsubstituted phenyl, monosubstituted phenyl or methylpyridinyl; (12) when Y' is  $CR^7R^7$ —C=O, W' is  $NR^9$ — $CR^7R^7$ ,  $Q_1$  and  $Q_2$  are hydrogen,  $A_1$  and  $A_2$  are CH, and  $Z_1$  and  $Z_2$  are O, then G-L is not unsubstituted phenyl; (13) when Y' is CHR<sup>7</sup>—NR<sup>7</sup> where R<sup>7</sup> is unsubstituted phenyl, methoxy or 25 ethoxy and R7 is unsubstituted phenyl, methyl or -C(O)-C6H5, W' is dimethoxyphenylene or unsubstituted phenylene,  $Z_1$  and  $Z_2$  are  $O,\,Q_1$  and  $Q_2$ are hydrogen, A<sub>1</sub> and A<sub>2</sub> are CH, C-CN, C-C(O)-C<sub>6</sub>H<sub>5</sub>, or -C(O)dimethoxyphenyl, then G-L is not unsubstituted phenyl; (14) the compound of formula Ia is not 6,10-epithio-4H-thieno-30

[3',4':5,6]cyclooct[1,2-f]isoindole-7,9(5H,8H)dione, 8-(3,5-dichlorophenyl)-

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6,6a,9a,10,11,12,-hexahydro-1,3,6,10-tetramethyl-2,2,13-trioxide, (6R,6aR,9aS,10S);

- (15) when Y' is O, W' is  $-CH_2-CH_2$ -,  $Q_1$  and  $Q_2$  are methyl,  $Z_1$  and  $Z_2$  are O, and  $A_1$  and  $A_2$  are CH, then G-L is not unsubstituted phenyl, phenyl substituted with methoxy, phenyl-alkyl-, or morpholine-alkyl, nor is the compound bridged to itself through a group L which is alkylene to form a bis compound;
- (16) when Y' is -O-,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is  $CR^7R^7$   $CR^7R^7$ , and  $A_1$  and  $A_2$  are CH, then G-L is not an unsubstituted phenyl group; and
- (17) when Y' is -O-,  $Q_1$  and  $Q_2$  are hydrogen,  $Z_1$  and  $Z_2$  are O, W' is cyclopentyl, cyclohexyl, 3-phenyl-2-isoxazoline or  $CR^7R^7$ - $CR^7R^7$  where  $R^7$  and  $R^7$  are each independently defined as Cl, Br, H and 4-butyrolactone and  $R^7$  and  $R^7$  are not all simultaneously H, and  $A_1$  and  $A_2$  are CH, then G-L is not an unsubstituted naphthyl ring or a monosubstituted phenyl ring, where said substituent is methoxy, Br, Cl,  $NO_2$ , methyl, ethyl,  $CH_2$ -phenyl, S-phenyl, or O-phenyl.

#### 2. The compound of Claim 1 wherein

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O, S, NH, or NR<sup>6</sup>;

 $Z_2$  is O, S, NH, or NR<sup>6</sup>;

 $A_1$  is  $CR^7$  or N;

25  $A_2$  is  $CR^7$  or N;

Y' is J-J'-J" where J is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, CR<sup>7</sup>R<sup>7'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, or heterocyclo or substituted heterocyclo, and J" is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, where Y' is not a bond;

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- W' is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein, when W' is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N;
- Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, or NR⁴R⁵;
- Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl,

  15 heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, or NR⁴R⁵;

L is a bond,  $(CR^7R^{7'})n$ , NH,  $NR^5$  or  $NR^5(CR^7R^{7'})n$ , where n = 0-3;

- 20 R¹ and R¹' are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
  - R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

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R³ and R³′ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, hydroxylamine, hydroxamide, alkoxy or substituted alkylthio;

R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

20 R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro,

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hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SOR<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

R<sup>8</sup> and R<sup>8'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR<sup>1</sup>, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, alkylthio or substituted alkylthio, C=OSR<sup>1</sup>, R<sup>1</sup>OC=O, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, R<sup>1</sup>R<sup>1'</sup>NC=O, S=OR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

with the provisos (1) to (17) of said formula Ia, and further where (i) when Y' is -O- and W' is  $CR^7R^7$ — $CR^7R^7$ ,  $A_1$  and  $A_2$  are not simultaneously CH; and (ii) when L is a bond, G is not an unsubstituted phenyl group.

# 3. The compound of Claim 1, wherein

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O;

 $Z_2$  is O;

 $A_1$  is  $CR^7$ ;

 $A_2$  is  $CR^7$ ;

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- Y' is J-J'-J'' where J is  $(CR^7R^{7'})n$  and n=0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH,  $NR^7$ ,  $CR^7R^{7'}$ ,  $R^2P=O$ ,  $R^2P=S$ ,  $R^2OP=O$ ,  $R^2NHP=O$ ,  $OP=OOR^2$ ,  $OP=ONHR^2$ ,  $OSO_2$ , NHNH,  $NHNR^6$ ,  $NR^6NH$ , N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, or heterocyclo or substituted heterocyclo, and J'' is  $(CR^7R^{7'})n$  and n=0-3, where Y' is not a bond;
- W' is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein, when W' is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N;
- Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R<sup>4</sup>C=O, R<sup>5</sup>R<sup>6</sup>NC=O, HOCR<sup>7</sup>R<sup>7'</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, or NR<sup>4</sup>R<sup>5</sup>;
- Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R<sup>4</sup>C=O, R<sup>5</sup>R<sup>6</sup>NC=O, HOCR<sup>7</sup>R<sup>7'</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, or NR<sup>4</sup>R<sup>5</sup>;

L is a bond;

R¹ and R¹ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

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- R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R³ and R³′ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, alkoxy or substituted alkoxy, amino, NR¹R², alkylthio or substituted alkylthio;
- R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
- R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
- R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O,
   R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

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R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR¹, amino, NHR⁴, NR²R⁵, alkylthio or substituted alkylthio, R¹C=O, R¹NHC=O, R¹R¹′NC=O, SO₂R¹, or SO₂NR¹R¹′; and

R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

with the provisos (1) to (17) of said formula Ia, and further where (i) when Y' is -O- and W' is  $CR^7R^{7'}$ — $CR^7R^{7'}$ ,  $A_1$  and  $A_2$  are not simultaneously CH; and (ii) when L is a bond, G is not an unsubstituted phenyl group.

4. A compound selected from the group consisting of:  $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Bromo-3-methylphenyl) tetrahydro-4,7-ethanothiopyrano[3,4-c]pyrrole-1,3,8(2H,4H)-trione (1C);$ 

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 $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Bromo-3-methylphenyl) tetrahydro-4,7-ethanothiopyrano[3,4-c]pyrrole-1,3,8(2H,4H)-trione 5,5-dioxide (2);\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(3-Chlorophenyl)hexahydro-4-methyl-4,\\ 7-epoxy-1H-isoindole-1,3(2H)-dione (3);$ 

- $\begin{array}{ll} 5 & (3a\alpha,4\alpha,7\alpha,7a\alpha)\text{- and }(3a\alpha,4\beta,7\beta,7a\alpha)\text{-}4\text{-}[(\text{Acetyloxy})\text{methyl}]\text{-}3a,4,7,7a-\\ & \text{tetrahydro-2-[3-(trifluoromethyl)phenyl}]\text{-}4,7\text{-}epoxy\text{-}1\text{H-isoindole-1,3(2H)-dione}\\ & (4i \& 4ii, \, \text{respectively});\\ & (3a\alpha,4\alpha,7\alpha,7a\alpha)\text{- and }(3a\alpha,4\beta,7\beta,7a\alpha)\text{-}4\text{-}[(\text{Acetyloxy})\text{methyl}]\text{-}Hexahydro-2\text{-}[3-(trifluoromethyl)phenyl}]\text{-}4,7\text{-}epoxy\text{-}1\text{H-isoindole-1,3(2H)-dione} \, (5i \& 5ii, )). \end{array}$
- 10 respectively);

 $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ - and  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -3a,4,7,7a-Tetrahydro-5-(hydroxymethyl)-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione (6i & 6ii, respectively);

 $(3a\alpha, 4\alpha, 7\alpha, 7a\alpha)$ -3a,4,7,7a-Tetrahydro-5-(hydroxymethyl)-4-methyl-2-[3-

- (trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione (7);
  (3aα,4β,7β,7aα)-2-[3,5-Bis(trifluoromethyl)phenyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione (8);
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Bromophenyl) octahydro-1,3-dioxo-4,7-etheno-5H-pyrrolo[3,4-c] pyridine-5-carboxylic acid phenyl ester (9);$
- $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Bromophenyl) octahydro-1,3-dioxo-4,7-etheno-5H-pyrrolo[3,4-c]pyridine-5-carboxylic acid phenylmethyl ester (10); \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-Hexahydro-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione trifluoroacetate (11); \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-5-Acetylhexahydro-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1$
- 25 1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione (12);  $(3a\alpha,4\alpha,7\alpha,7a\alpha)\text{-}5\text{-}Benzoylhexahydro-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione (13); <math display="block">(3a\alpha,4\alpha,7\alpha,7a\alpha)\text{-}Hexahydro-5\text{-}methyl-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione (14);$

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- $(3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{Hexahydro-5-(phenylmethyl)-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione trifluoroacetate (15); \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{Hexahydro-5-propyl-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-1H-pyrrolo[3,4-c]pyridine-1,3(2H)-dione trifluoroacetate (16); \\$
- 5 (3aα,4α,4aβ,5aβ,6α,6aα)-2-[4-Cyano-3-(trifluoromethyl)phenyl]decahydro-1,3-dioxo-4,6-(iminomethano)cycloprop[f]isoindole-7-carboxylic acid phenylmethyl ester (17);
  - $(3a\alpha,4\alpha,4a\beta,5a\beta,6\alpha,6a\alpha)-4-[Decahydro-1,3-dioxo-4,6-(iminomethano)cycloprop[f]isoindol-2-yl]-2-(trifluoromethyl)benzonitrile (18);$
- 10  $(3a\alpha,4\alpha,4a\beta,5a\beta,6\alpha,6a\alpha)$ -4-[Decahydro-7-methyl-1,3-dioxo-4,6-(iminomethano)cycloprop[f]isoindol-2-yl]-2-(trifluoromethyl)benzonitrile (19);  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile (20B);
- (3aα,4β,7β,7aα)-N-[4-[[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethyl]thio]phenyl]acetamide
- (21E);  $(3a\alpha,4\beta,7\beta,7a\alpha)\text{-N-[4-[[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-}$

methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethyl]sulfinyl]phenyl]acetamide (22);

- 20  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -N-[4-[[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethyl]sulfonyl]phenyl]acetamide (23);
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$  and  $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -N-[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-
- yl]ethyl]benzenesulfonamide (24Ci & 24Cii, respectively);  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-(2-hydroxyethyl)-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile (25B); <math display="block">(3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{ and }(3a\alpha,4\beta,7\beta,7a\alpha)-N-[4-[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-$
- 30 yl]ethoxy]phenyl]acetamide (26Ci & 26Cii, respectively);

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 $(3a\alpha, 4\alpha, 7\alpha, 7a\alpha)$ -Hexahydro-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione (27D);

 $(1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta,6a\alpha)$ -Hexahydro-4-(2-naphthalenyl)-2,6-epoxy-3Hoxireno[f]isoindole-3,5(4H)-dione (28B);

- $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-[4-Bromo-3-(trifluoromethyl)phenyl]-3a,4,7,7a-tetrahydro-4,7-dimethyl-4,7-epithio-1H-isoindole-1,3(2H)-dione 8-oxide (29); \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-2-[4-Bromo-3-(trifluoromethyl)phenyl]-3a,4,7,7a-tetrahydro-4,7-epithio-1H-isoindole-1,3(2H)-dione 8-oxide (30); \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-Hexahydro-2-[3-(trifluoromethyl)phenyl]-4,7-imino-1H-$
- isoindole-1,3(2H)-dione (31D);  $(3a\alpha,4\beta,7\beta,7a\alpha)-\text{ and }(3a\alpha,4\alpha,7\alpha,7a\alpha)-3a,4,7,7a-\text{Tetrahydro-4,7-dimethyl-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione (32i & 32ii, respectively);}$

(3a $\alpha$ ,4 $\alpha$ ,7 $\alpha$ ,7a  $\alpha$ )-Hexahydro-4,7-dimethyl-2-[3-(trifluoromethyl)phenyl]-4,7-

- epoxy-1H-isoindole-1,3(2H)-dione (33);  $(3a\alpha,4\alpha,7\alpha,7a\alpha)\text{-Tetrahydro-5-methyl-2-(4-nitro-1-naphthalenyl)-4,7-etheno-1H-pyrrolo[3,4-c]pyridine-1,3,6(2H,5H)-trione (34B); \\ (3a\alpha,4\beta,7\beta,7a\alpha)\text{-4-[4-[2-(4-Fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile (35);}$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-(2-Bromoethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile (36); \\ (3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4,7-dimethyl-2-(3-methyl-4-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione (37); \\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(2-Fluorenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-$
- dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)-2-[3-Chloro-4-(4-morpholinyl)phenyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(2,3-Dihydro-1H-inden-5-yl)hexahydro-4,7-epoxy-$

isoindole-1,3(2H)-dione;

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- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-1-naphthalenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Chloro-1-naphthalenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(5-Amino-1-naphthalenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $3a\alpha,4\beta,7\beta,7a\alpha$ )-Hexahydro-2-(7-hydroxy-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-
- 10 isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(1H-indol-5-yl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(1H-indazol-6-yl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(1,3-Benzodioxol-5-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[4-Amino-3-(trifluoromethyl)phenyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Chloro-4-iodophenyl)hexahydro-4,7-epoxy-1H-
- 20 isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(8-quinolinyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(2,3-Dihydro-1,4-benzodioxin-6-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[2-oxo-4-(trifluoromethyl)-2H-1-benzopyran-7-yl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(4-methyl-2-oxo-2H-1-benzopyran-7-yl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(2,5-Dimethoxy-4-nitrophenyl)hexahydro-4,7-epoxy-1H-
- 30 isoindole-1,3(2H)-dione;

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- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2,3,5,6-Tetrafluoro-4-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2,4,5-trifluorophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2,4,5-trichlorophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(2-Amino-4,5-dichlorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,4-Difluorophenyl)hexahydro-4,7-epoxy-1H-isoindole-
- 10 1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -1-Acetyl-2,3-dihydro-6-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-1H-indole;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Chloro-4-fluorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,4-Dichlorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(3,4,5-trichlorophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Chloro-4-methoxyphenyl)hexahydro-4,7-epoxy-1H-
- 20 isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Chloro-4-methylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-methyl-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Chloro-3-methylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,4-Dimethylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[4-Bromo-3-(trifluoromethyl)phenyl]hexahydro-4,7-epoxy-
- 30 1H-isoindole-1,3(2H)-dione;

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 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-3-methylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Fluoro-3-nitrophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[4-Fluoro-3-(trifluoromethyl)phenyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Chloro-3-nitrophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[4-Chloro-3-(trifluoromethyl)phenyl]hexahydro-4,7-epoxy-
- 10 1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Chloro-2-methoxy-5-methylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Amino-3-nitrophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15 (3aα,4β,7β,7aα)-Hexahydro-2-(4-methyl-3-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,4-Dimethoxyphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(3-hydroxy-4-methoxyphenyl)-4,7-epoxy-1H-20 isoindole-1,3(2H)-dione;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(4-methyl-5-nitro-2-pyridinyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-Chloro-4-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)- $\alpha$ -phenylbenzeneacetonitrile;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-methoxy-3-dibenzofuranyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2,3,4-trifluorophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

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 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(2,3-Dihydro-2-methyl-1,3-dioxo-1H-isoindol-5-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-2,3,5,6-tetrafluorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-hydroxy-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[2,5-Dichloro-4-(1H-pyrrol-1-yl)phenyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[4-(methoxymethyl)-2-oxo-2H-1-benzopyran-7-
- 10 yl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione; (3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )-2-(6-Benzothiazolyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-Methoxy-4-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzoic acid methyl ester;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-Methyl-5-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-oxo-2H-1-benzopyran-6-yl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 20 1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2,4,5-trimethylphenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Fluoro-3-methylphenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(3-methoxy-4-methylphenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -N-Ethyl-2-methyl-5-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-N-phenylbenzenesulfonamide;
  - $(3a\alpha, 4\beta, 7\beta, 7a\alpha) 2, 6 Dibromo 4 (octahydro 1, 3 dioxo 4, 7 epoxy 2H isoindol 1, 3 dioxo 1,$
- 30 2-yl)benzenesulfonamide;

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 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2,4-Dimethyl-6-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-3-pyridinecarbonitrile;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(2,3-Dimethyl-1H-indol-5-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Dibenzofuranyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2'-hydroxy[1,1':3',1"-terphenyl]-5'-yl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(5,6,7,8-tetrahydro-3-hydroxy-2-naphthalenyl)-
- 4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(2,3-Dihydro-1H-indol-6-yl)hexahydro-4,7-epoxy-1H-indol-6-yl)hexahydro-6-yl$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(1,3-Dihydro-2,2-dioxidobenzo[c]thiophen-5-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-hydroxy-4,5-dimethylphenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(2,3-Dihydro-2,2,3,3-tetrafluoro-1,4-benzodioxin-6-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(1H-indazol-5-yl)-4,7-epoxy-1H-isoindole-
- 20 1,3(2H)-dione;

isoindole-1,3(2H)-dione;

- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Amino-2,3,5,6-tetrafluorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-3-chlorophenyl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(5-hydroxy-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-(Octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Morpholinyl)-5-(octahydro-1,3-dioxo-4,7-epoxy-2H-
- 30 isoindol-2-yl)benzoic acid methyl ester;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-Fluoro-5-(octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

- (3aα,4β,7β,7aα)-2-(9-Ethyl-9H-carbazol-3-yl)hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-2-[1,2-Dihydro-8-methyl-2-oxo-4-(trifluoromethyl)-7-quinolinyl]hexahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -Hexahydro-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-
- isoindole-1,3(2H)-dione;  $(3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{Hexahydro-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-3-methylphenyl)-3a,4,7,7 $\alpha$ -tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- (3aα,4β,7β,7aα)-3a,4,7,7a-Tetrahydro-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-2-(9-Ethyl-9H-carbazol-3-yl)-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-2-[4-Fluoro-3-(trifluoromethyl)phenyl]-3a,4,7,7a-tetrahydro-
- 4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)-2-[1,2-Dihydro-8-methyl-2-oxo-4-(trifluoromethyl)-7-quinolinyl]-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione; <math display="block">(3a\alpha,4\alpha,7\alpha,7a\alpha)-4-[(Acetyloxy)methyl]-2-(4-bromo-3-methylphenyl)-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[(Acetyloxy)methyl]-2-(4-bromo-3-methylphenyl)-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione.;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4,7-dimethyl-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-1,3)-4-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-dimethyl-1,$
- 30 2-yl)-1-naphthalenecarbonitrile;

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1H-isoindole-1,3(2H)-dione;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -(Benzo[b]thiophen-3-yl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4,7-dimethyl-2-[4-nitro-3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-(1,3,3a,4,7,7a-Hexahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-1-naphthalenecarbonitrile;\\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-Hexahydro-4-methyl-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Bromo-3-methylphenyl)hexahydro-4-methyl-4,7-epoxy-
- 10 1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)\text{-Hexahydro-4-methyl-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-}$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,5-Dichlorophenyl)hexahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3-Chloro-4-fluorophenyl)hexahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-Methoxy-4-(octahydro-1,3-dioxo-4-methyl-4,7-epoxy-2H-isoindol-2-yl)-1-naphthalenecarbonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4-methyl-2-[4-nitro-3-(trifluoromethyl)phenyl]-4,7-20 epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[4-(1H-imidazol-1-yl)phenyl]-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
    - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[3-Chloro-4-(2-thiazolyl)phenyl]hexahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25  $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -2-(3,5-Dichlorophenyl)hexahydro-4,7-imino-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -2-(4-Bromo-1-naphthalenyl)hexahydro-4,7-imino-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -2-(4-Bromo-3-methylphenyl)hexahydro-4,7-imino-1H-
- 30 isoindole-1,3(2H)-dione;

- $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -Hexahydro-2-(4-nitro-1-naphthalenyl)-4,7-imino-1H-isoindole-1,3(2H)-dione;
- $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -8-Acetyl-2-(3,5-dichlorophenyl)hexahydro-4,7-imino-1H-isoindole-1,3(2H)-dione;
- (3aα,4α,7α,7aα)-Octahydro-1,3-dioxo-2-[3-(trifluoromethyl)phenyl]-4,7-ethano-5H-pyrrolo[3,4-c]pyridine-5-carboxylic acid phenyl ester;
   (3aα,4α,7α,7aα)-4-(Octahydro-1,3-dioxo-4,7-ethano-2H-pyrrolo[3,4-c]pyridin-2-yl)-1-naphthalenecarbonitrile;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -4-(Octahydro-5-methyl-1,3-dioxo-4,7-ethano-2H-pyrrolo[3,4-
- c]pyridin-2-yl)-1-naphthalenecarbonitrile;
   (3aα,4α,7α,7aα)-2-(4-Cyano-1-naphthalenyl)octahydro-1,3-dioxo-4,7-etheno-5H-pyrrolo[3,4-c]pyridine-5-carboxylic acid phenylmethyl ester;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -4-(Octahydro-1,3-dioxo-4,7-ethano-2H-pyrrolo[3,4-c]pyridin-2-yl)-2-(trifluoromethyl)benzonitrile;
- (3aα,4α,7α,7aα)-4-(Octahydro-5-methyl-1,3-dioxo-4,7-ethano-2H-pyrrolo[3,4-c]pyridin-2-yl)-2-(trifluoromethyl)benzonitrile;
   (3aα,4α,7α,7aα)-2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-1,3-dioxo-4,7-etheno-5H-pyrrolo[3,4-c]pyridine-5-carboxylic acid phenylmethyl ester;

 $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-[4-Bromo-3-(trifluoromethyl)phenyl]tetrahydro-5-methyl-$ 

- 4,7-etheno-1H-pyrrolo[3,4-c]pyridine-1,3,6(2H,5H)-trione;  $(3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{Tetrahydro-5-methyl-2-[3-(trifluoromethyl)phenyl]-4,7-etheno-1H-pyrrolo[3,4-c]pyridine-1,3,6(2H,5H)-trione; \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-\text{Tetrahydro-5-methyl-2-(2-naphthalenyl)-4,7-etheno-1H-pyrrolo[3,4-c]pyridine-1,3,6(2H,5H)-trione; }$
- $(1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta,6a\alpha)-Hexahydro-4-[3-(trifluoromethyl)phenyl]-2,6-epoxy-3H-oxireno[f]isoindole-3,5(4H)-dione;\\ (1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta,6a\alpha)-4-(3,5-Dichlorophenyl)hexahydro-2,6-epoxy-3H-oxireno[f]isoindole-3,5(4H)-dione;\\ (1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta,6a\alpha)-Hexahydro-4-(4-nitro-1-naphthalenyl)-2,6-epoxy-$
- 30 3H-oxireno[f]isoindole-3,5(4H)-dione;

- $(1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta,6a\alpha)$ -4-(3,4-Dichlorophenyl)hexahydro-2,6-epoxy-3H-oxireno[f]isoindole-3,5(4H)-dione;
- 2-[4-(4-Bromophenoxy)phenyl]-3a,4,7,7a-tetrahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 5 3a,4,7,7a-Tetrahydro-2-(2-methoxyphenyl)-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-;
  - [(1,2,3,3a,7,7a-Hexahydro-2-phenyl-4,7-epoxy-4H-isoindol-4-yl)methyl]carbamic acid (3,5-dimethoxyphenyl)methyl ester;
  - 2-(2,4-Dimethylphenyl)-3a,4,7,7a-tetrahydro-4-(hydroxymethyl)-4,7-epoxy-1H-
- 10 isoindole-1,3(2H)-dione;
  - 2-(1,3-Benzodioxol-5-yl)-3a,4,7,7a-tetrahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 4-[Bis(acetyloxy)methyl]-2-(3-bromophenyl)-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- N-[[1,2,3,3a,7,7a-Hexahydro-2-(2,4,6-trimethylphenyl)-4,7-epoxy-4H-isoindol-4-yl]methyl]-2,2-dimethylpropanamide;
  - 3a,4,7,7a-Tetrahydro-4-(hydroxymethyl)-2-[2-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 3a,4,7,7a-Tetrahydro-4-(hydroxymethyl)-2-(1-naphthalenyl)-4,7-epoxy-1H-
- 20 isoindole-1,3(2H)-dione;
  - 2-Chloro-5-(1,3,3a,4,7,7a-hexahydro-4,7-dimethyl-4,7-epoxy-2H-isoindol-2-yl)benzoic acid methyl ester;
  - 4-[Bis(acetyloxy)methyl]-2-(4-bromo-2-nitrophenyl)-3a,4,7,7a-tetrahydro-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25 3a,4,7,7a-Tetrahydro-4-methyl-2-(4-methyl-3-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 2-[2-Chloro-5-(trifluoromethyl)phenyl]-3a,4,7,7a-tetrahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 2-[4-Chloro-3-(trifluoromethyl)phenyl]-3a,4,7,7a-tetrahydro-4,7-dimethyl-4,7-
- 30 epoxy-1H-isoindole-1,3(2H)-dione;
  - 2-(1,3,3a,4,7,7a-Hexahydro-4-methyl-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;

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- 2-(4-Fluorophenyl)-3a,4,7,7a-tetrahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 2,2,2-Trifluoro-N-[(1,2,3,3a,7,7a-hexahydro-2-phenyl-4,7-epoxy-4H-isoindol-4-yl)methyl]acetamide;
- 5 3a,4,7,7a-Tetrahydro-4,7-dimethyl-2-(4-methyl-3-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 2-Chloro-5-[1,3,3a,4,7,7a-hexahydro-4-(hydroxymethyl)-4,7-epoxy-2H-isoindol-2-yl]benzoic acid;
  - 3a,4,7,7a-Tetrahydro-4,7-dimethyl-2-(4-nitrophenyl)-4,7-epoxy-1H-isoindole-
- 10 1,3(2H)-dione;
  - 3a,4,7,7a-Tetrahydro-2-(2-mercaptophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - 3a,4,7,7a-Tetrahydro-2-[2-[(phenylmethyl)thio]phenyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15 [[2-(4-Chlorophenyl)-1,2,3,3a,7,7a-hexahydro-4,7-epoxy-4H-isoindol-4-yl]methyl]carbamic acid 2-methylpropyl ester;
  - 4-(1,1-Dimethylethyl)-N-[[1,2,3,3a,7,7a-hexahydro-2-(4-methylphenyl)-4,7-epoxy-4H-isoindol-4 yl]methyl]benzamide;
  - 2,4-Dichloro-N-[[1,2,3,3a,7,7a-hexahydro-2-(4-nitrophenyl)-4,7-epoxy-4H-
- 20 isoindol-4-yl]methyl]benzamide;
  - N-[[2-(4-Chlorophenyl)-1,2,3,3a,7,7a-hexahydro-4,7-epoxy-4H-isoindol-4-yl]methyl]-2,4,6-trimethylbenzenesulfonamide;
  - [(1,2,3,3a,7,7a-Hexahydro-2-phenyl-4,7-epoxy-4H-isoindol-4-yl)methyl]carbamic acid 1,1-dimethylethyl ester;
- N-[(1,2,3,3a,7,7a-Hexahydro-2-phenyl-4,7-epoxy-4H-isoindol-4-yl)methyl]-2-phenoxyacetamide;
  - N-[[1,2,3,3a,7,7a-Hexahydro-2-(4-nitrophenyl)-4,7-epoxy-4H-isoindol-4-yl]methyl]-2,2-dimethylpropanamide;
  - 2-(2,4-Dichlorophenoxy)-N-[[1,2,3,3a,7,7a-hexahydro-2-(4-nitrophenyl)-4,7-
- 30 epoxy-4H-isoindol-4-yl]methyl]acetamide;

- N-[[1,2,3,3a,7,7a-Hexahydro-2-(4-methylphenyl)-4,7-epoxy-4H-isoindol-4-yl]methyl]-3,5-dimethoxybenzamide;
- N-[[2-(4-Chlorophenyl)-1,2,3,3a,7,7a-hexahydro-4,7-epoxy-4H-isoindol-4-yl]methyl]-2-nitrobenzenesulfonamide;
- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[(1S)-1-phenylethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[(1S)-2-hydroxy-1-phenylethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-[(1S)-2-(Acetyloxy)-1-phenylethyl]-3a,4,7,7a-tetrahydro-
- 10 4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\alpha,7\alpha,7a\alpha)$ -3a,4,7,7a-Tetrahydro-2-[(1S)-1-phenylethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[(1R)-1-phenylethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 15 (3aα,4β,7β,7aα)-4-[[[(Octahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)methyl]amino]benzoic acid;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-(4-morpholinylmethyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-(2-hydroxyethyl)-7-methyl-1,3-dioxo-4,7-epoxy-
- 20 2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;(3aα,4β,7β,7aα)- and (3aα,4α,7α,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-
  - (phenylmethyl)-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -(4-[7-[2-(4-Bromophenoxy)ethyl]octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
- $25 \qquad (3a\alpha, 4\beta, 7\beta, 7a\alpha)-4-[Octahydro-7-[2-(4-iodophenoxy)ethyl]-4-methyl-1, 3-dioxo-4, 7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl) benzonitrile;$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;

- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-7-[2-(4-methoxyphenoxy)ethyl]-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[7-[2-(4-Ethoxyphenoxy)ethyl]octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\$
- (3aα,4β,7β,7aα)-4-[7-[2-(4-Chlorophenoxy)ethyl]octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
   (3aα,4β,7β,7aα)-4-[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethoxy]benzoic acid, methyl ester;
- (3aα,4β,7β,7aα)-Hexahydro-4-(2-hydroxyethyl)-7-methyl-2-(3-methyl-4-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethoxy)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(3,5-Dichlorophenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4,7-dimethyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-[4-Cyano-3-(trifluoromethyl)phenyl-1,3-epoxy-1H-isoindole-1,3(H)-$
- dioxo-4,7-epoxy-4H-isoindole-4-propanenitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[2-(4-morpholinyl)ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile, trifluoroacetate; <math display="block">(3a\alpha,4\beta,7\beta,7a\alpha)-2-(5-Fluoro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(5-Fluoro-4-nitro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(1,1-Dioxidobenzo[b]thiophen-3-yl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ 4-(1,3,3a,4,7,7a-Hexahydro-4,6,7-trimethyl-1,3-dioxo-4,7-epoxy-2H-isoindole-1,3(2H)-dioxo-4,7-epoxy-2H-isoindole-1,3(H)-dioxo-4,7-epoxy-2$
- 30 pyrrolo[3,4-c]pyridin-2-yl)-2-(trifluoromethyl)benzonitrile;

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- $(3a\alpha,4\beta,7\beta,7a\alpha)-Tetrahydro-4,7-dimethyl-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3,5(2H,4H)-trione;\\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-Tetrahydro-4,7-dimethyl-2-[3-(trifluoromethyl)phenyl]-4,7-epoxy-1H-isoindole-1,3,5(2H,4H)-trione;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(5-Chloro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(5-Chloro-4-nitro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-Ethylhexahydro-7-methyl-2-(4-nitro-1-naphthalenyl)-4,7-$
- epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl)-N-(4-fluorophenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetamide; <math display="block">(3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4-methyl-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione, faster eluting enantiomer;$
- (3aα,4β,7β,7aα)-Hexahydro-4-methyl-2-(2-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione, slower eluting enantiomer;
   (3aα,4β,7β,7aα)-4-[4-[2-[[(4-Fluorophenyl)methyl]methylamino]ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
- (3aα,4β,5β,6β,7β,7aα)-4-(Octahydro-4,5,6,7-tetramethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;
   (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile, faster eluting antipode;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile, slower eluting enantiomer;  $(3a\alpha,4\beta,5\beta,7\beta,7a\alpha)$ -4-(Octahydro-5-hydroxy-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;

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 $(3a\alpha,4\beta,5\alpha,7\beta,7a\alpha)-4-(Octahydro-5-hydroxy-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile; \\ (\alpha R)-\alpha-Methoxybenzeneacetic acid, 2-[(3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl$ 

- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(Methylthio)-4-(octahydro-4,7-dimethyl-1,3-dioxo-4,7-dimethyl-1,3-dioxo-4,7-dimethyl-1,3-dioxo-4,7-dimethyl-2-yl) benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(Methylsulfinyl)-4-(octahydro-4,7-dimethyl-1,3-dioxo-4,7-dime$
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(Methylsulfinyl)-4-(octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(Methylsulfonyl)-4-(octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;\\ (3a\alpha,4\beta,5\beta,7\beta,7a\alpha)-7-[2-[[(1,1-Dimethylethyl)dimethylsilyl]oxy]ethyl]hexahydro-5-hydroxy-4-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- $(3a\alpha,4\beta,5\beta,7\beta,7a\alpha)-\text{Hexahydro-5-hydroxy-7-(2-hydroxyethyl)-4-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}\\ (3a\alpha,4\beta,5\beta,7\beta,7a\alpha)-7-[2-(4-Fluorophenoxy)ethyl]\text{hexahydro-5-hydroxy-4-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}\\ (3a\alpha,4\beta,5\beta,6\beta,7\beta,7a\alpha)-4-(Octahydro-5,6-dihydroxy-4,7-dimethyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1,3-dioxo-1-methyl-1-methyl-1,3-dioxo-1-methyl-1-methyl-1,3-dioxo-1-methyl-1-methyl-1,3-dioxo-1-methyl-1-methyl-1,3-dioxo-1-methyl-1-methyl-1-methyl-1-methyl-1-methyl-1-methyl-1-methyl-1-methyl-1-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1-methyl-1$
- 4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;  $(3a\alpha,4\beta,5\alpha,6\alpha,7\beta,7a\alpha)-4-(Octahydro-5,6-dihydroxy-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile; <math display="block">3a\alpha,4\beta,5\beta,6\beta,7\beta,7a\alpha)-4-[Octahydro-5,6-dihydroxy-4-(hydroxyethyl)-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- $(3a\alpha,4\beta,5\beta,6\beta,7\beta,7a\alpha)-4-[Octahydro-5,6-dihydroxy-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2- (trifluoromethyl)benzonitrile;$ 
  - $(3a\alpha,4\beta,5\beta,5a\beta,8a\beta,8b\alpha)-4-(Decahydro-5-hydroxy-4-methyl-1,3-dioxo-4,8a-epoxy-2H-furo[3,2-e]isoindol-2-yl)-1-naphthalenecarbonitrile;$

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetic acid;

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetic acid, methyl ester;

- 5  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Cyano-1-naphthalenyl)-N-[(4-fluorophenyl)methyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetamide;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-N-[2-[2-(4-Cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethyl]-4-fluorobenzamide;$
- [3aR-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[Octahydro-4-(2-hydroxyethyl)-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[Octahydro-4-(2-hydroxyethyl)-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- 1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aS-(-(3aα,4β,7β,7aα)]-4-[4-[2-(3-Fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; (4-Fluorophenyl)carbamic acid, 2-[(3aα,4β,7β,7aα)-2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl
- 20 ester;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-(2-hydroxyethyl)-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ (3a\alpha,4\beta,6\beta,7\beta,7a\alpha)-4-[4-[2-(4-Cyanophenoxy)ethyl]octahydro-6-hydroxy-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- 25 [3aS-(3aα,4β,5β,7β,7aα)]-4-[Octahydro-5-hydroxy-7-(2-hydroxyethyl)-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aR-(3aα,4β,5β,7β,7aα)]-4-[Octahydro-5-hydroxy-7-(2-hydroxyethyl)-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; (3aα,4β,7β,7aα)-4-[4-[2-(4-Cyanophenoxy)ethyl]-7-ethyloctahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- 30 4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

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- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[2-(Acetyloxy)ethyl]-2-(4-cyano-1-naphthalenyl)hexahydro-7-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-methyl-1,3-dioxo-7-(2-oxoethyl)-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- $[3a\alpha,4\beta(E),7\beta,7a\alpha]-4-[4-[3-(4-Cyanophenyl)-2-propenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3a\alpha,4\beta(Z),7\beta,7a\alpha]-4-[4-[3-(4-Cyanophenyl)-2-propenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[3-(4-Cyanophenyl)propyl]octahydro-7-methyl-1,3-$
- dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-[(6-Chloro-1,2-benzisoxazol-3-yl)oxy]ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; <math display="block">(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[2-[(6-nitro-1H-indazol-3-yl)oxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- [3aS-(3a $\alpha$ ,4 $\beta$ ,5 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[7-[2-(1,2-Benzisoxazol-3-yloxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aR-(3a $\alpha$ ,4 $\beta$ ,5 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[7-[2-(1,2-Benzisoxazol-3-yloxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-
- 20 naphthalenecarbonitrile;  $(3a\alpha,4\beta,5\beta,7\beta,7a\alpha)]\text{-}4\text{-}(Octahydro-5\text{-}hydroxy-4,7\text{-}dimethyl-1,3\text{-}dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;}\\ (3a\alpha,4\beta,5\beta,7\beta,7a\alpha)]\text{-}4\text{-}(Octahydro-5\text{-}hydroxy-4,7\text{-}dimethyl-1,3\text{-}dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethyl)benzonitrile;}$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl) octahydro-1,3-dioxo-7-[2-(phenylmethoxy)ethyl]-4,7-epoxy-4H-isoindole-4-propanenitrile;\\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Cyano-1-naphthalenyl) octahydro-1,3-dioxo-7-[2-(phenylmethoxy)ethyl]-4,7-epoxy-4H-isoindole-4-propanenitrile;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl) octahydro-7-(2-hydroxyethyl)-$
- 30 1,3-dioxo-4,7-epoxy-4H-isoindole-4-propanenitrile;

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 $(3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(4-Cyano-1-naphthalenyl) octahydro-7-(2-hydroxyethyl)-1,3-dioxo-4,7-epoxy-4H-isoindole-4-propanenitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl)-7-[2-(4-fluorophenoxy)ethyl] octahydro-1,3-dioxo-4,7-epoxy-4H-isoindole-4-propanenitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl)-7-[2-(4-Cyano-1-naphthalenyl)-7$ 

- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(7-Chloro-2,1,3-benzoxadiazol-4-yl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(7-Chloro-2-methyl-4-benzofuranyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- (3aα,4β,7β,7aα)-2-(7-Chloro-2-methylbenzo[b]thiophen-4-yl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   [3aα,4β(E),7β,7aα]-4-[2-(4-Cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]-2-butenoic acid, phenylmethyl ester;
   (3aα,4β,7β,7aα)-2-(4-Cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-
- 4,7-epoxy-4H-isoindole-4-butanoic acid;
   (3aα,4β,7β,7aα)-2-(4-Cyano-1-naphthalenyl)-N-(4-fluorophenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-butanamide;
   [3aS-(3aα,4β,5β,7β,7aα)]-4-[7-[2-(Acetyloxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- [3aR-(3a $\alpha$ ,4 $\beta$ ,5 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[Octahydro-5-hydroxy-7-(2-hydroxyethyl)-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; (3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ (E)]-4-[Octahydro-4-methyl-1,3-dioxo-7-(4-oxo-4-phenyl-2-butenyl)-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; (3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ (E)]-4-[Octahydro-4-methyl-1,3-dioxo-7-(4-oxo-4-phenyl-2-
- butenyl)-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)\text{-}(4\text{-}[7\text{-}[2\text{-}(4\text{-Bromophenoxy})\text{ethyl}]\text{octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;} \\ (3a\alpha,4\beta,7\beta,7a\alpha)\text{-}4\text{-}[\text{Octahydro-7-}[2\text{-}(4\text{-iodophenoxy})\text{ethyl}]\text{-}4\text{-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl}]\text{-}2-(trifluoromethyl)benzonitrile;}$

- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-7-[2-(4-methoxyphenoxy)ethyl]-4-methyl-1,3-
- dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[7-[2-(4-Ethoxyphenoxy)ethyl]octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[7-[2-(4-Chlorophenoxy)ethyl]octahydro-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;$
- 10 (3aα,4β,7β,7aα)-4-[2-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethoxy]benzoic acid, methyl ester;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4-(2-hydroxyethyl)-7-methyl-2-(3-methyl-4-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethoxy)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
   (3aα,4β,7β,7aα)-2-(3,5-Dichlorophenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- (3aα,4β,7β,7aα)-Hexahydro-4,7-dimethyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(phenylmethoxy)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4-(2-hydroxyethyl)-7-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[2-(4-Fluorophenoxy)ethyl]hexahydro-7-methyl-2-(3-methyl-4-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

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 $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-[(trifluoromethyl)thio]phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;$ 

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-methyl-7-[2-(4-nitrophenoxy)ethyl]-1,3-dioxo-

- 5 4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[2-(4-Fluorophenoxy)ethyl] hexahydro-7-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
  - (3aα,4β,7β,7aα)-4-[Octahydro-7-methyl-1,3-dioxo-7-[2-[2-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-
- 10 (trifluoromethyl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(2-Bromophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(3-Fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-2-[4-(1H-imidazol-1-yl)phenyl]-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-[3-Chloro-4-(2-thiazolyl)phenyl]hexahydro-4-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$ 
  - (3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )-Hexahydro-4,7-dimethyl-2-(3-methyl-4-nitrophenyl)-4,7-
- 20 epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4,7-dimethyl-2-(2-methyl-4-nitrophenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(3,5-Dichlorophenyl)hexahydro-4-(2-hydroxyethyl)-7-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- 25 (3aα,4β,7β,7aα)-2-(3,5-Dichlorophenyl)-4-[2-(4-fluorophenoxy)ethyl]hexahydro-7-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[Octahydro-4-[2-(4-hydroxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;

 $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(4-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;\\(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[3-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2-$ 

- 5 (trifluoromethyl)benzonitrile;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(3-Bromophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[(4-Fluorophenyl)methyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\$
- (3aα,4β,7β,7aα)-2-(1,6-Dihydro-1-methyl-6-oxo-3-pyridinyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-Hexahydro-4,7-dimethyl-2-(1-methyl-6-oxo-3-piperidinyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-4-[4-[2-(3-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-
- dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethoxy]benzoic acid, phenylmethyl ester; <math display="block">(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-1,3-dioxo-7-(2-phenoxyethyl)-4,7-$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(3,5-Dichloro-4-nitrophenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(3,5-Dichloro-4-hydroxyphenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$

epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile;

- $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(5-Fluoro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-25 epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4,7-dimethyl-2-(1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[3-methoxy-4-(5-oxazolyl)phenyl]-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

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- $(3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4-[2-(4-methoxyphenoxy)ethyl]-7-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-Hexahydro-4-methyl-2-(4-nitro-1-naphthalenyl)-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-\text{Hexahydro-4-methyl-2-(4-nitro-1-naphthalenyl)-7-[2-(4-nitrophenoxy)ethyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(1,6-Dihydro-1,4-dimethyl-6-oxo-3-pyridinyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-7-methyl-2-(4-nitro-1-naphthalenyl)-1,3-dioxo-1-methyl-2$
- 4,7-epoxy-4H-isoindol-4-yl]ethoxy]benzonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-1,2-benzenedicarbonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-(2-Bromoethyl)hexahydro-7-methyl-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(4-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-[2-(4-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-$
- dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(3-Fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; <math display="block">(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[2-[3-(4-morpholinyl)phenoxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[2-[4-nitro-3-(triflouoromethyl)phenoxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[4-[2-(3-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\$

- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-(2,3-Dihydro-3-methyl-2-oxo-6-benzothiazolyl) hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-(2,3-Dihydro-2-oxo-6-benzothiazolyl) hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;$
- 2H-isoindol-2-yl)phenyl]acetamide;
   (3aα,4β,7β,7aα)-4-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)-2-(trifluoromethoxy)benzonitrile;
   (3aα,4β,7β,7aα)-2-Methoxy-4-(octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)benzonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)-2-[4-(4,5-Dichloro-1H-imidazol-1-yl)phenyl]hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-[4-(4-Bromo-1-methyl-1H-pyrazol-3-yl)phenyl]hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-(2-hydroxyethyl)-7-methyl-1,3-dioxo-4,7-dioxo-4,$
- 20 epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)-2\text{-lodo-4-}(Octahydro-4,7\text{-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl)}benzonitrile; \\ (3a\alpha,4\beta,7\beta,7a\alpha)-4\text{-[4-[2-(4-Fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;}$
- (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
   (3aα,4β,7β,7aα)-4-[4-[2-(4-Cyano-3-fluorophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

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 $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-[2,3,5,6-tetrafluoro-4-(trifluoromethyl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$ 

 $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4,7-dimethyl-2-[4-(1H-1,2,4-triazol-3-yl)phenyl]-

- $\label{eq:continuous} 4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\ (3a\alpha,4\beta,7\beta,7a\alpha)-2-[4-(4,5-Dihydro-5-oxo-1,2,4-oxadiazol-3-yl)phenyl]hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\$ 
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-2-[3-methoxy-4-(2-oxazolyl)phenyl]-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
- (3aα,4β,7β,7aα)-Hexahydro-2-(4-hydroxy-1-naphthalenyl)-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-Hexahydro-2-(8-hydroxy-5-quinolinyl)-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione, trifluoroacetate;

(3aα.4β.7β.7aα)-4-[Octahydro-4-methyl-1,3-dioxo-7-[2-

- [methyl(phenylmethyl)amino]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-2- (trifluoromethyl)benzonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)\text{-Hexahydro-4,7-dimethyl-2-(5-quinolinyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione;} \\ (3a\alpha,4\beta,7\beta,7a\alpha)\text{-5-(Octahydro-4,7-dimethyl-1,3-dioxo-4,7-epoxy-2H-isoindol-1)}$
- 20 2-yl)-2-pyridinecarbonitrile;  $(3a\alpha,4\beta,7\beta,7a\alpha)\text{-}5\text{-}(Octahydro-4,7\text{-}dimethyl-1,3\text{-}dioxo-4,7\text{-}epoxy-2H-isoindol-2-yl)-8-quinolinecarbonitrile;} \\ (3a\alpha,4\beta,7\beta,7a\alpha)\text{-}2\text{-}(5\text{-}Bromo-4\text{-}nitro-1\text{-}naphthalenyl)hexahydro-4,7\text{-}dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;}$
- 25  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(5-Bromo-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -Hexahydro-4,7-dimethyl-2-[8-(trifluoromethyl)-4-quinolinyl]-4,7-epoxy-1H-isoindole-1,3(2H)-dione;

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- 4-Fluorobenzoic acid, 2-[ $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
- Benzeneacetic acid, 2-[  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-cyano-1-
- 5 naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
  - 4-Fluorobenzeneacetic acid, 2-[  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
- ( $3a\alpha,4\beta,7\beta,7a\alpha$ )-Hexahydro-4-methyl-7-[2-[4-(methylsulfonyl)phenoxy]ethyl]-2-(4-nitro-1-naphthalenyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione; ( $3a\alpha,4\beta,7\beta,7a\alpha$ )-Hexahydro-2-(2-naphthalenyl)-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-Chloro-1-naphthalenyl)hexahydro-4,7-dimethyl-4,7-
- epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-N-[(4-Chlorophenyl)methyl]-2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetamide;
  - 4,7,7-Trimethyl-3-oxo-2-oxabicyclo[2.2.1]heptane-1-carboxylic acid, 2-
- 20 [ $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
  - ( $\alpha$ S)-  $\alpha$ -Methoxy- $\alpha$ -(trifluoromethyl)benzeneacetic acid, 2-[( $3a\alpha$ , $4\beta$ , $7\beta$ , $7a\alpha$ )-2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
- 25  $(\alpha R)$   $\alpha$ -Methoxy- $\alpha$ -(trifluoromethyl)benzeneacetic acid, 2-[( $3a\alpha$ , $4\beta$ , $7\beta$ , $7a\alpha$ )-2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
  - $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[2-[(7-methyl-1,2-benzisoxazol-3-yl)oxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- 30  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[4-[2-(1,2-Benzisoxazol-3-yloxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

1H-isoindole-1,3(2H)-dione;

5

dione;

 $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[2-(Benzoyloxy)ethyl]-2-(4-cyano-1-naphthalenyl)hexahydro-7-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;\\(3a\alpha,4\beta,7\beta,7a\alpha)-2-(4-Cyano-1-naphthalenyl)-4-[2-[(4-nitrobenzoyl)oxy]ethyl]hexahydro-7-methyl-4,7-epoxy-1H-isoindole-1,3(2H)-1,3($ 

- 4-Chlorobenzoic acid, 2-[ $(3a\alpha,4\beta,7\beta,7a\alpha)$ -2-(4-cyano-1-naphthalenyl)octahydro-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-y]ethyl ester;
- [3aα,4β,7β,7aα(E)]-4-[Octahydro-4-methyl-7-[3-(1-naphthalenyl)-2-propenyl] 1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
   (3aα,4β,7β,7aα)-4-[Octahydro-4-methyl-7-[3-(1-naphthalenyl)propyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
   (3aα,4β,7β,7aα)-Hexahydro-4,7-dimethyl-2-(2-methyl-6-quinolinyl)-4,7-epoxy-
- (3aα,4β,7β,7aα)-Hexahydro-2-(5-isoquinolinyl)-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-2-(6-Benzothiazolyl)hexahydro-4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   [3aα,4β,7β,7aα(E)]-4-[Octahydro-4-methyl-1,3-dioxo-7-(4-oxo-4-phenyl-2-
- butenyl)-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
   (3aα,4β,7β,7aα)-2-(4-Cyano-1-naphthalenyl)octahydro-N-(2-hydroxyphenyl)-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindole-4-acetamide;
   [3aα,4β(E),7β,7aα]-4-[Octahydro-4-methyl-7-[3-(6-methyl-2-pyridinyl)-2-propenyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- $(3a\alpha,4\beta,7\beta,7a\alpha)-4-[Octahydro-4-methyl-7-[3-(6-methyl-2-pyridinyl)propyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;\\[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-4-[2-(3-methoxyphenoxy)ethyl]-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1,3-dioxo-4,3-$
- 30 1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

[3aR-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[4-[2-(4-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[4-[2-(4-Cyanophenoxy)ethyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

- $(3a\alpha,4\alpha,7\alpha,7a\alpha)-4-[4-[(4-Fluorophenyl)methyl]octahydro-7-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-2-(trifluoromethyl)benzonitrile; \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-Hexahydro-4,7-dimethyl-2-(1-methyl-6-oxo-3-piperidinyl)-4,7-epoxy-1H-isoindole-1,3(2H)-dione; \\ (3a\alpha,4\alpha,7\alpha,7a\alpha)-2-(1,6-Dihydro-1,4-dimethyl-6-oxo-3-pyridinyl)hexahydro-$
- 4,7-dimethyl-4,7-epoxy-1H-isoindole-1,3(2H)-dione;
   (3aα,4β,7β,7aα)-2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-1,3-dioxo-7-[2-(phenylmethoxy)ethyl]-4,7-epoxy-4H-isoindole-4-propanenitrile;
   (3aα,4β,7β,7aα)-2-[4-Cyano-3-(trifluoromethyl)phenyl]octahydro-1,3-dioxo-7-[2-(phenylmethoxy)ethyl]-4,7-epoxy-4H-isoindole-4-propanenitrile;
- 15  $(3a\alpha,4\beta,7\beta,7a\alpha)$ -4-[7-[2-(4-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[7-[2-(1,3-Benzodioxol-5-yloxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(1,3-Benzodioxol-5-yloxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-[(5-Chloro-2-pyridinyl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-$
- 25 naphthalenecarbonitrile;  $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-[(5-Chloro-2-pyridinyl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; <math display="block">[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(4-Chlorophenoxy)ethyl]octahydro-5-hydroxy-4-30 methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$

 $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(4-Chlorophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(4-Acetylphenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\$ 

- $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(4-Acetylphenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; \\ [3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3-Cyanophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-4-methyl-1,3-dioxo-4,7-epoxy-4-methyl-1,3-dioxo-4,7-epoxy-2-methy$
- methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-5-hydroxy-4-methyl-1,3-dioxo-7-[2-[(5,6,7,8-tetrahydro-1-naphthalenyl)oxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$ 
  - $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[Octahydro-5-hydroxy-4-methyl-1,3-dioxo-7-[2-methyl-1,3-dioxo-7-[2-methyl-1,3-dioxo-7-[2-methyl-1,3-dioxo-7-[2-methyl-1,3-dioxo-7-[3-methyl-1,3-methyl$
- 15 [(5,6,7,8-tetrahydro-1-naphthalenyl)oxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
  - [3aS- $(3a\alpha,4\beta,7\beta,7a\alpha)$ ]-4-[Octahydro-5-hydroxy-4-methyl-1,3-dioxo-7-[2-[(5,6,7,8-tetrahydro-5-oxo-1-naphthalenyl)oxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
- 20 [3aR-(3aα,4β,7β,7aα)]-4-[Octahydro-5-hydroxy-4-methyl-1,3-dioxo-7-[2-[(5,6,7,8-tetrahydro-5-oxo-1-naphthalenyl)oxy]ethyl]-4,7-epoxy-2H-isoindol-2yl]-1-naphthalenecarbonitrile;
  - $[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(4-Fluorophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- [3aR- $(3a\alpha,4\beta,7\beta,7a\alpha)$ ]-4-[7-[2-(4-Fluorophenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; [3aS- $(3a\alpha,4\beta,7\beta,7a\alpha)$ ]-4-[Octahydro-5-hydroxy-4-methyl-7-[2-[(4-methyl-2-oxo-2H-1-benzopyran-7-yl)oxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

[ $3aR-(3a\alpha,4\beta,7\beta,7a\alpha)$ ]-4-[Octahydro-5-hydroxy-4-methyl-7-[2-[(4-methyl-2-oxo-2H-1-benzopyran-7-yl)oxy]ethyl]-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

[3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[7-[2-(3,5-Dimethoxyphenoxy)ethyl]octahydro-5-

- 5 hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
  - $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-(3,5-Dimethoxyphenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;$
- [3aR-(3aα,4β,7β,7aα)]-]-4-[7-[2-(4-Chloro-3-methylphenoxy)ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
  [3aR-(3aα,4β,7β,7aα)]-4-[7-[2-(4-Cyano-2,3-difluorophenoxy)ethyl]octahydro-

5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-

- naphthalenecarbonitrile;  $[3aS-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-[(5-Chloro-1,2-benzisoxazol-3-yl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; <math display="block">[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-[(5-Chloro-1,2-benzisoxazol-3-yl]-4-[7-[2-[(5-Chloro-1,2-$
- yl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;  $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-3-[2-[2-(4-Cyano-1-naphthalenyl)octahydro-6-hydroxy-7-methyl-1,3-dioxo-4,7-epoxy-4H-isoindol-4-yl]ethoxy]-5-isoxazolecarboxylic acid, methyl ester;$
- [3aR-(3aα,4β,7β,7aα)]-4-[Octahydro-5-hydroxy-4-methyl-1,3-dioxo-7-[2-[4-(1H-1,2,4-triazol-1-yl)phenoxy]ethyl]-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;
   [3aS-(3aα,4β,7β,7aα)]-4-[7-[2-[(7-Chloro-4-quinolinyl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile, trifluoroacetate;

 $[3aR-(3a\alpha,4\beta,7\beta,7a\alpha)]-4-[7-[2-[(7-Chloro-4-quinolinyl)oxy]ethyl]octahydro-5-hydroxy-4-methyl-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile, trifluoroacetate;$ 

 $(1a\alpha,2\beta,2a\alpha,5a\alpha,6\beta b,6a\alpha)$ -4-[2-[2-[[(1,1-dimethylethyl)-

dimethylsilyl]oxy]ethyl]octahydro- 6-methyl-3,5-dioxo-2,6-epoxy-4H-oxireno[f]isoindol-4-yl]-1-naphthalenecarbonitrile;

[3aR- $(3a\alpha,4\beta,7\beta,7a\alpha)$ ]-4-[4-Ethyloctahydro-7-(2-hydroxyethyl)-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile;

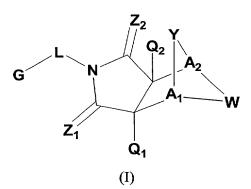
[3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[4-Ethyloctahydro-7-(2-hydroxyethyl)-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitril;

[3aR-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[4-[2-(4-Cyanophenoxy)ethyl]-7-ethyloctahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile; and [3aS-(3a $\alpha$ ,4 $\beta$ ,7 $\beta$ ,7a $\alpha$ )]-4-[4-[2-(4-Cyanophenoxy)ethyl]-7-ethyloctahydro-1,3-dioxo-4,7-epoxy-2H-isoindol-2-yl]-1-naphthalenecarbonitrile.

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5. A pharmaceutical composition capable of treating a NHR - associated condition, comprising a compound of the following formula I or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier:



20

wherein the symbols have the following meanings and are, for each occurrence, independently selected:

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

1.4

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LD0192 (NP)
       Express Mail Number EL596928345US
       Z_1 is O, S, NH, or NR^6;
       Z_2 is O, S, NH, or NR<sup>6</sup>;
       A_1 is CR^7 or N;
       A_2 is CR^7 or N;
       Y is J-J'-J" where J is (CR^7R^{7'})n and n = 0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH,
5
                 NR<sup>7</sup>, C=O, OC=O, NR<sup>1</sup>C=O, CR<sup>7</sup>R<sup>7'</sup>, C=CR<sup>8</sup>R<sup>8'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O,
                 R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OP=OR<sup>2</sup>, OSO<sub>2</sub>, C=NR<sup>7</sup>, NHNH,
                 NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or
                  substituted cycloalkenyl, heterocyclo or substituted heterocyclo or aryl or
                  substituted aryl, and J" is (CR^7R^{7'})n and n = 0-3, where Y is not a bond;
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        W is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>8</sup>=CR<sup>8'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—
                  NR<sup>9'</sup>, S—CR<sup>7</sup>R<sup>7'</sup>, SO—CR<sup>7</sup>R<sup>7'</sup>, SO<sub>2</sub>—CR<sup>7</sup>R<sup>7'</sup>, cycloalkyl or substituted
                  cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or
                  substituted heterocyclo, or aryl or substituted aryl, wherein when W is not
                  NR^9—CR^7R^{7'}, N=CR^8, N=N, NR^9—NR^{9'}, S—CR^7R^{7'}, SO—CR^7R^{7'}, SO
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                  CR<sup>7</sup>R<sup>7'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O,
                  SO<sub>2</sub>, NH, NR<sup>7</sup>, OC=O, NR<sup>1</sup>C=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH,
                  NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N;
         Q1 is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or
                   substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl,
                   heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted
                   arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo
                   or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R6NC=O,
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- 20 HOCR<sup>7</sup>R<sup>7'</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup> or NR<sup>4</sup>R<sup>5</sup>;
- Q2 is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or 25 substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R6NC=O,

HOCR<sup>7</sup>R<sup>7'</sup>, nitro, R<sup>1</sup>OCH<sub>2</sub>, R<sup>1</sup>O, NH<sub>2</sub>, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup> or NR<sup>4</sup>R<sup>5</sup>; 30

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L is a bond,  $(CR^7R^{7'})n$ , NH, NR<sup>5</sup>, NH $(CR^7R^{7'})n$  or NR<sup>5</sup> $(CR^7R^{7'})n$ , where n = 0-3;

R<sup>1</sup> and R<sup>1'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

R³ and R³′ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, hydroxylamine, hydroxamide, alkoxy or substituted alkoxy, amino, NR¹R², thiol, alkylthio or substituted alkylthio;

20 R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>;

R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

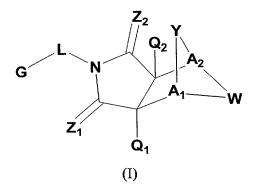
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- R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR1, R1C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>';
- R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR1, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>, PO<sub>2</sub>R<sup>1</sup>R<sup>1</sup>, R<sup>1</sup>R<sup>1</sup>NC=O<sub>2</sub>C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>, or, wherein 15  $A_1$ , or  $A_2$  contains a group  $R^7$  and W contains a group  $R^7$ , said  $R^7$  groups of  $A_1$ or A2 and W together form a heterocyclic ring;
- R<sup>8</sup> and R<sup>8'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or 20 substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR1, amino, NHR4, NR2R5, NOR<sup>1</sup>, alkylthio or substituted alkylthio, C=OSR<sup>1</sup>, R<sup>1</sup>OC=O, R<sup>1</sup>C=O,  $R^{1}NHC=O,\,R^{1}R^{1'}NC=O,\,SO_{2}OR^{1},\,S=OR^{1},\,SO_{2}R^{1},\,PO_{3}R^{1}R^{1'},\,or\,SO_{2}NR^{1}R^{1'};$ 25 and
  - R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or

substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR $^1$ , R $^1$ C=O, R $^1$ OC=O, R $^1$ NHC=O, SO $_2$ R $^1$ , SO $_2$ OR $^1$ , or SO $_2$ NR $^1$ R $^1$ '.

- 6. A pharmaceutical composition of Claim 5 further comprising anotheranti-cancer agent.
  - 7. A method of modulating the function of a nuclear hormone receptor which comprises administering to a mammalian species in need thereof an effective nuclear hormone receptor modulating amount of a compound of the following formula I:



wherein the symbols have the following meanings and are, for each occurrence, independently selected:

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O, S, NH, or NR<sup>6</sup>;

20  $Z_2$  is O, S, NH, or NR<sup>6</sup>;

A<sub>1</sub> is CR<sup>7</sup> or N;

A<sub>2</sub> is CR<sup>7</sup> or N;

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Y is J-J'-J" where J is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, C=O, OC=O, NR<sup>1</sup>C=O, CR<sup>7</sup>R<sup>7'</sup>, C=CR<sup>8</sup>R<sup>8'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OP=OR<sup>2</sup>, OSO<sub>2</sub>, C=NR<sup>7</sup>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or

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substituted aryl, and J" is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, where Y is not a bond;
W is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>8</sup>=CR<sup>8'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, S—CR<sup>7</sup>R<sup>7'</sup>, SO—CR<sup>7</sup>R<sup>7'</sup>, SO<sub>2</sub>—CR<sup>7</sup>R<sup>7'</sup>, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein when W is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, S—CR<sup>7</sup>R<sup>7'</sup>, SO—CR<sup>7</sup>R<sup>7'</sup>, SO<sub>2</sub>—CR<sup>7</sup>R<sup>7'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O,

SO<sub>2</sub>, NH, NR<sup>7</sup>, OC=O, NR<sup>1</sup>C=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH,

substituted cycloalkenyl, heterocyclo or substituted heterocyclo or aryl or

10 NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N;

Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR³R⁵′, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴R⁵;

Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR³R⁵¹, nitro, R¹OCH₃, R¹O, NH₃, C=OSR¹, SO₃R¹ or NR⁴R⁵;

L is a bond,  $(CR^7R^{7'})n$ , NH, NR<sup>5</sup>, NH $(CR^7R^{7'})n$  or NR<sup>5</sup> $(CR^7R^{7'})n$ , where n = 0-3;

R¹ and R¹′ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

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- R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R³ and R³ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, hydroxylamine, hydroxamide, alkoxy or substituted alkylthio;
- R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>':
- R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>;
- R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O,
  R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>;

- R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkylalkyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, heterocycloalkylalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>, or, wherein A<sub>1</sub> or A<sub>2</sub> contains a group R<sup>7</sup> and W contains a group R<sup>7</sup>, said R<sup>7</sup> groups of A<sub>1</sub> or A<sub>2</sub> and W together form a heterocyclic ring;
- R<sup>8</sup> and R<sup>8'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR<sup>1</sup>, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, alkylthio or substituted alkylthio, C=OSR<sup>1</sup>, R<sup>1</sup>OC=O, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, R<sup>1</sup>R<sup>1'</sup>NC=O, SO<sub>2</sub>OR<sup>1</sup>, S=OR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>; and
- R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>.
- 8. The method of Claim 7 wherein said nuclear hormone receptor is a30 steroid binding nuclear hormone receptor.

- 9. The method of Claim 7 wherein said nuclear hormone receptor is the androgen receptor.
- 10. The method of Claim 7 wherein said nuclear hormone receptor is theestrogen receptor.
  - 11. The method of Claim 7 wherein said nuclear hormone receptor is the progesterone receptor.
- 10 12. The method of Claim 7 wherein said nuclear hormone receptor is the glucocorticoid receptor.
  - 13. The method of Claim 7 wherein said nuclear hormone receptor is the mineralocorticoid receptor.
  - 14. The method of Claim 7 wherein said nuclear hormone receptor is the aldosterone receptor.
- 15. A method for treating a condition or disorder comprising administering20 to a mammalian species in need thereof a therapeutically effective amount of a compound of the following formula:

$$\begin{array}{c|c}
Z_2 & Y \\
Q_2 & A_2 \\
\hline
Z_1 & Q_1
\end{array}$$
(I)

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wherein the symbols have the following meanings and are, for each occurrence, independently selected:

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

5  $Z_1$  is O, S, NH, or NR<sup>6</sup>;

 $Z_2$  is O, S, NH, or NR<sup>6</sup>;

 $A_1$  is  $CR^7$  or N;

 $A_2$  is  $CR^7$  or N;

Y is J-J'-J'' where J is  $(CR^7R^{7'})$ n and n = 0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH,

NR<sup>7</sup>, C=O, OC=O, NR<sup>1</sup>C=O, CR<sup>7</sup>R<sup>7'</sup>, C=CR<sup>8</sup>R<sup>8'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OP=OR<sup>2</sup>, OSO<sub>2</sub>, C=NR<sup>7</sup>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo or aryl or substituted aryl, and J" is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, where Y is not a bond;

W is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>, CR<sup>8</sup>=CR<sup>8'</sup>, CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, S—CR<sup>7</sup>R<sup>7'</sup>, SO—CR<sup>7</sup>R<sup>7'</sup>, SO<sub>2</sub>—CR<sup>7</sup>R<sup>7'</sup>, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein when W is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, S—CR<sup>7</sup>R<sup>7'</sup>, SO—CR<sup>7</sup>R<sup>7'</sup>, SO<sub>2</sub>—

20 CR<sup>7</sup>R<sup>7'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OC=O, NR<sup>1</sup>C=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N;

 $Q_1$  is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl,

heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴R⁵;

Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl,

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heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴R⁵;

- L is a bond, (CR<sup>7</sup>R<sup>7'</sup>)n, NH, NR<sup>5</sup>, NH(CR<sup>7</sup>R<sup>7'</sup>)n or NR<sup>5</sup>(CR<sup>7</sup>R<sup>7'</sup>)n, where n = 0-3;

  R<sup>1</sup> and R<sup>1'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R² is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, hydroxylamine, hydroxamide, alkoxy or substituted alkoxy, amino, NR<sup>1</sup>R<sup>2</sup>, thiol, alkylthio or substituted alkylthio;

R<sup>3</sup> and R<sup>3'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or

- R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl
  or substituted cycloalkenyl, heterocyclo or substituted heterocyclo,
  cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted
  cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or
  substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O,
  SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
- 30 R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo,

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cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>.  $SO_2OR^1$ , or  $SO_2NR^1R^{1'}$ :

5 R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, 10  $R^1NHC=O$ ,  $SO_2R^1$ ,  $SO_2OR^1$ , or  $SO_2NR^1R^{1'}$ ;

R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>,

PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>, or, wherein  $A_1$  or  $A_2$  contains a group  $R^7$  and W contains a group  $R^7$ , said  $R^7$  groups of  $A_1$ or A<sub>2</sub> and W together form a heterocyclic ring;

 $R^8$  and  $R^{8'}$  are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or 25 substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, nitro, halo, CN, OR<sup>1</sup>, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, alkylthio or substituted alkylthio, C=OSR<sup>1</sup>, R<sup>1</sup>OC=O, R<sup>1</sup>C=O,  $R^{1}NHC=O$ ,  $R^{1}R^{1}NC=O$ ,  $SO_{2}OR^{1}$ ,  $S=OR^{1}$ ,  $SO_{3}R^{1}$ ,  $PO_{3}R^{1}R^{1}$ , or  $SO_{3}NR^{1}R^{1}$ ;

30 and

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R<sup>9</sup> and R<sup>9'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

wherein said condition or disorder is selected from the group consisting of proliferate diseases, cancers, benign prostate hypertrophia, adenomas and neoplasies of the prostate, benign or malignant tumor cells containing the androgen receptor, heart disease, angiogenic conditions or disorders, hirsutism, acne, hyperpilosity, inflammation, immune modulation, seborrhea, endometriosis, polycystic ovary syndrome, androgenic alopecia, hypogonadism, osteoporosis, suppressing spermatogenisis, libido, cachexia, anorexia, inhibition of muscular atrophy in ambulatory patients, androgen supplementation for age related decreased testosterone levels in men, cancers expressing the estrogen receptor. prostate cancer, breast cancer, endometrial cancer, hot flushes, vaginal dryness, menopause, amennoreahea, dysmennoreahea, contraception, pregnancy termination, cancers containing the progesterone receptor, endometriosis, cachexia, menopause, cyclesynchrony, meniginoma, fibroids, labor induction, autoimmune diseases, Alzheimer's disease, psychotic disorders, drug dependence, non-insulin dependent Diabetes Mellitus, dopamine receptor mediated disorders, congestive heart failure, disregulation of cholesterol homeostasis, and attenuating the metabolism of a pharmaceutical agent.

16. A method for preparation of a compound of the following formula XVI, or salt thereof:

where

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G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O, S, NH, or NR<sup>6</sup>;

 $Z_2$  is O, S, NH, or NR<sup>6</sup>;

 $A_1$  is  $CR^7$  or N;

 $A_2$  is  $CR^7$  or N;

10 Y' is J-J'-J" where J is  $(CR^7R^{7'})n$  and n = 0-3, J' is O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OC=O, NR<sup>1</sup>C=O, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N, and J" is  $(CR^7R^{7'})n$  and n = 0-3;

Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴Rҕ;

Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R⊓, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴Rҕ;

25 L is a bond,  $(CR^7R^{7'})n$ , NH,  $NR^5$  or  $NR^5(CR^7R^{7'})n$ , where n = 0-3;

R<sup>1</sup> and R<sup>1'</sup> are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl,

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cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;

- R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
- R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>,
   SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;
  - R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>; and
  - R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or

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substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

5 comprising the steps of contacting a compound of the following formula XV, or salt thereof:

$$\begin{array}{c|c}
Z_2 & Y \\
\hline
Z_1 & Q_2 \\
Z_1 & Q_1
\end{array}$$
XV

- where the symbols are as defined above; with an enzyme or microorganism capable of catalyzing the hydroxylation of said compound XV to said compound XVI, and effecting said hydroxylation.
- 17. A method for preparation of a compound of the following formula 15 XVIII, or salt thereof:

$$\begin{array}{c|c}
 & Z_2 & Y \\
 & Q_2 \\
 & Q_2 \\
 & Q_2 \\
 & Q_1 \\
 & Q_1 \\
 & Q_1
\end{array}$$
OH

where

G is an aryl or heterocyclo group, where said group is mono- or polycyclic, and which is optionally substituted at one or more positions;

 $Z_1$  is O, S, NH, or NR<sup>6</sup>;

 $Z_2$  is O, S, NH, or NR<sup>6</sup>;

 $A_1$  is  $CR^7$  or N;

25  $A_2$  is  $CR^7$  or N;

Y' is J-J'-J" where J is  $(CR^7R^{7'})n$  and n = 0-3, J' is O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OC=O, NR<sup>1</sup>C=O, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N, and J" is  $(CR^7R^{7'})n$  and n = 0-3;

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- Q<sub>1</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R¹′, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴R⁵;
- Q<sub>2</sub> is H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocycloalkyl or substituted heterocycloalkyl, arylalkyl or substituted arylalkyl, alkynyl or substituted alkynyl, aryl or substituted aryl, heterocyclo or substituted heterocyclo, halo, CN, R¹OC=O, R⁴C=O, R⁵R⁶NC=O, HOCR⊓R¹, nitro, R¹OCH₂, R¹O, NH₂, C=OSR¹, SO₂R¹ or NR⁴R⁵;

L is a bond,  $(CR^7R^{7'})n$ , NH,  $NR^5$  or  $NR^5(CR^7R^{7'})n$ , where n = 0-3;

- R¹ and R¹′ are each independently H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkyalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- 20 R<sup>2</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl;
- 25 R<sup>4</sup> is H, alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O,

  SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1</sup>';

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R<sup>5</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, eycloalkenylalkyl or substituted cycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

R<sup>6</sup> is alkyl or substituted alkyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkenylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, CN, OH, OR<sup>1</sup>, R<sup>1</sup>C=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>; and

R<sup>7</sup> and R<sup>7'</sup> are each independently H, alkyl or substituted alkyl, alkenyl or substituted alkenyl, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, cycloalkylalkyl or substituted cycloalkylalkyl, cycloalkenylalkyl or substituted cycloalkylalkyl, heterocycloalkyl or substituted heterocycloalkyl, aryl or substituted aryl, arylalkyl or substituted arylalkyl, halo, CN, OR<sup>1</sup>, nitro, hydroxylamine, hydroxylamide, amino, NHR<sup>4</sup>, NR<sup>2</sup>R<sup>5</sup>, NOR<sup>1</sup>, thiol, alkylthio or substituted alkylthio, R<sup>1</sup>C=O, R<sup>1</sup>OC=O, R<sup>1</sup>NHC=O, SO<sub>2</sub>R<sup>1</sup>, SOR<sup>1</sup>, PO<sub>3</sub>R<sup>1</sup>R<sup>1'</sup>, R<sup>1</sup>R<sup>1'</sup>NC=O, C=OSR<sup>1</sup>, SO<sub>2</sub>R<sup>1</sup>, SO<sub>2</sub>OR<sup>1</sup>, or SO<sub>2</sub>NR<sup>1</sup>R<sup>1'</sup>;

comprising the steps of contacting a compound of the following formula XVII, or salt thereof:

$$\begin{array}{c|c}
Z_2 & Y \\
Q_2 & A_2 \\
Z_1 & Q_1
\end{array}$$

XVII

where the symbols are as defined above; with an enzyme or microorganism capable of catalyzing the opening of the epoxide ring of compound XVII to form the diol of said compound XVIII, and effecting said ring opening and diol formation.

## 18. A compound of the following formula Ib:

$$\begin{array}{c|c}
Z_2 \\
Q_2 \\
A_1 \\
Q_1
\end{array}$$
Ib

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where  $G, Z_1, Z_2, Q_1$  and  $Q_2$  are as defined in claim 1;

Y' is J-J'-J" where J is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, J' is a bond or O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, CR<sup>7</sup>R<sup>7'</sup>, R<sup>2</sup>P=O, R<sup>2</sup>P=S, R<sup>2</sup>OP=O, R<sup>2</sup>NHP=O, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, N=N, cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, or heterocyclo or substituted heterocyclo, and J" is (CR<sup>7</sup>R<sup>7'</sup>)n and n = 0-3, where Y is not a bond; and W' is CR<sup>7</sup>R<sup>7'</sup>—CR<sup>7</sup>R<sup>7'</sup>—C=O, NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>,

cycloalkyl or substituted cycloalkyl, cycloalkenyl or substituted cycloalkenyl, heterocyclo or substituted heterocyclo, or aryl or substituted aryl, wherein,

when W' is not NR<sup>9</sup>—CR<sup>7</sup>R<sup>7'</sup>, N=CR<sup>8</sup>, N=N, NR<sup>9</sup>—NR<sup>9'</sup>, or heterocyclo or substituted heterocyclo, then J' must be O, S, S=O, SO<sub>2</sub>, NH, NR<sup>7</sup>, OP=OOR<sup>2</sup>, OP=ONHR<sup>2</sup>, OSO<sub>2</sub>, NHNH, NHNR<sup>6</sup>, NR<sup>6</sup>NH, or N=N; or alternatively,

Y' is  $CR^7R^{7'}$ -C=O and W' is  $NR^9$ - $CR^7R^{7'}$ ;

20 L is a bond; and

 $A_1$  and  $A_2$  are as defined above with the proviso that, when Y' = O and  $W' = -CH_2$ - $CH_2$ -, then at least one of  $A_1$  or  $A_2$  is not CH;
with the further provisos (2), (3), (6), (7) and (8) of claim 1.